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Application No. 08/356,229 Attorney Docket No.06/82-50439

an alkyl or aromatic organic [a] spacer molecule bound to the binding group [surface]; and

a carbohydrate derivative, O-, N-, C-, or S-glycosidically bound to the spacer molecule, which carbohydrate derivative specifically binds in a sample to at least one member selected from the group consisting of a protein, a virus and a cell.

23. (Amended) The biosensor according to claim 22, wherein said carbohydrate derivative is a [biologically active part] fragment of a naturally occurring carbohydrate sequence, which fragment binds in a biospecific manner to at least one member selected from the group consisting of a protein, a virus and a cell.

24. (Amended) The biosensor according to claim 22, wherein said <u>binding group</u> [spacer] is chemically bound or is bound via adsorption to the surface of the biosensor.

25. The biosensor according to claim 22, wherein said surface comprises a signal transducer.

26. The biosensor according to claim 22, wherein said surface comprises a means for monitoring a physical signal.



21. The biosensor according to claim 26, wherein said means for monitoring a physical signal is at least one member selected from the group consisting of a photometer, a chemical electrode, an electrochemical electrode, a temperature signal transducer, and a pressure signal transducer.

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3 28. (Amended) The biosensor according to claim 23, wherein the fragment of a naturally occurring carbohydrate sequence [said biologically active carbohydrate derivative] is a member selected from the group consisting of a mono-, di-, tri-, tetra-, or penta-saccharide sequence.

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29. (Twice Amended) The biosensor according to claim 23, wherein the fragment of a naturally occurring carbohydrate sequence [said biologically active carbohydrate derivative] selectively binds to at least one member selected from the group consisting of a lectin, an antibody against a carbohydrate, a cancer cell, a protein associated with a blood group determinant, a pathogenic bacteria, a pathogenic virus, a pathogenic toxin, a protein associated with an inflammatory reaction, and a cell associated with an inflammatory reaction.

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36. (Amended) The biosensor according to claim 23, wherein the fragment of a naturally occurring carbohydrate sequence [said carbohydrate derivative] binds to P-fimbriated E. coli.

The biosensor according to claim 22, wherein said carbohydrate derivative comprises at least one component selected from the group consisting of hexosamine-, fucose-, galactose-, glucose-, mannose-, xylose-, a N-acetylneuraminic acid residue, and analogs thereof.

The biosensor according to claim 31, wherein the carbohydrate derivative has been derivatized in at least one hydroxyl group or amino group thereof with an organic or inorganic group.

23. The biosensor according to claim 22, in which the carbohydrate derivative contains at least one O-, N-, S-, or C-glycosidically bound aglycon.

34. The biosensor according to claim 33, in which the aglycon contains at least one aliphatic or aromatic compound.

carbohydrate derivative contains an amino acid <u>component</u> [-], peptide <u>component</u> [-], or protein **molecule**• component.

26. (Amended) The biosensor according to claim 22, in which the carbohydrate derivative comprises at least one of a glycoprotein and [or] a neoglycoprotein.

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The biosensor according to claim 22, wherein said surface is operably associated with an optical sensor which gives a signal change upon binding of a protein, a virus or a cell to the carbohydrate derivative bound via the spacer to the surface.

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The biosensor according to claim 37, wherein the optical sensor functions by at least one method selected from the group consisting of surface plasmon changes, ellipsometry, reflection measurement and polarization measurement.

39. The biosensor according to claim 22, in which the surface is operably associated with a member selected from the group consisting of a piezoelectric crystal, an electrochemical electrode and a thermistor.

40. The biosensor according to claim 22, wherein said surface of the biosensor comprises gold.

Amended) A method of using the biosensor according to claim 22 to determine the presence or amount of a protein, a virus or a cell, comprising the steps of:

exposing the biosensor to a sample containing a protein, a virus or a cell to be measured,

binding a protein, virus or cell to the biosensor, <u>and</u>
measuring the presence or amount of the protein, virus or cell in the sample.

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[44], further comprising a protein which is linked between the spacer molecule and the <u>binding</u> group [biosensor surface].

46. The biosensor according to claim 22, wherein said spacer comprises albumin.

47. The biosensor according to claim 22, wherein said spacer comprises a protein.

The biosensor according to claim 25, wherein said signal transducer is a chemical transducer.

The biosensor according to claim 25, wherein said signal transducer is a physical transducer.

Please add new claims 50-84, as follows:

The immobilized carbohydrate derivative biosensor according to claim 22, further comprising a protein which is linked between the binding group and the biosensor surface.

The biosensor according to claim 46, wherein said carbohydrate derivative is a fragment of a naturally occurring carbohydrate sequence, which fragment binds in a biospecific manner to at least one member selected from the group consisting of a protein, a virus and a cell.





The biosensor according to claim 50, wherein said carbohydrate derivative is a fragment of a naturally occurring carbohydrate sequence, which fragment binds in a biospecific manner to at least one member selected from the group consisting of a protein, a virus and a cell.

53. The biosensor according to claim 48, wherein said surface comprises a signal transducer.

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54. The biosensor according to claim 50, wherein said surface comprises a signal transducer.

58. The biosensor according to claim 45, wherein said surface comprises a means for monitoring a physical signal.

56. The biosensor according to claim 50, wherein said surface comprises a means for monitoring a physical signal.

57. The biosensor according to claim 55, wherein said means for monitoring a physical signal is at least one member selected from the group consisting of a photometer, a chemical electrode, an electrochemical electrode, a temperature signal transducer, and a pressure signal transducer.

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58. The biosensor according to claim 56, wherein said means for monitoring a physical signal is at least one member selected from the group consisting of a photometer, a chemical electrode, an electrochemical electrode, a temperature signal transducer, and a pressure signal transducer.

59. The biosensor according to claim 51, wherein the fragment of a naturally occurring carbohydrate sequence is a member selected from the group consisting of a mono-, di-, tri-, tetra-, or penta-saccharide sequence.

60. The biosensor according to claim 52, wherein the fragment of a naturally occurring carbohydrate sequence is a member selected from the group consisting of a mono-, di-, tri-, tetra-, or penta-saccharide sequence.

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64. The biosensor according to claim 54, wherein the fragment of a naturally occurring carbohydrate sequence selectively binds to at least one member selected from the group consisting of a lectin, a cancer cell, a protein associated with a blood group determinant, a pathogenic bacteria, a pathogenic virus, a pathogenic toxin, a protein associated with an inflammatory reaction, and a cell associated with an inflammatory reaction.

62. The biosensor according to claim 52, wherein the fragment of a naturally occurring

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carbohydrate sequence selectively binds to at least one member selected from the group consisting of a lectin, a cancer cell, a protein associated with a blood group determinant, a pathogenic bacteria, a pathogenic virus, a pathogenic toxin, a protein associated with an inflammatory reaction, and a cell associated with an inflammatory reaction.

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63. The biosensor according to claim 45, wherein said surface is operably associated with an optical sensor which gives a signal change upon binding of a protein, a virus or a cell to the carbohydrate derivative.

64. The biosensor according to claim 56, wherein said surface is operably associated with an optical sensor which gives a signal change upon binding of a protein, a virus or a cell to the carbohydrate derivative.

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65. The biosensor according to claim 63, wherein the optical sensor functions by at least one method selected from the group consisting of surface plasmon changes, ellipsometry, reflection measurement and polarization measurement.

66. The biosensor according to claim 64, wherein the optical sensor functions by at least one method selected from the group consisting of surface plasmon changes, ellipsometry, reflection measurement and polarization measurement.

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The biosensor according to claim A5, in which the surface is operably associated with a member selected from the group consisting of a piezoelectric crystal, an electrochemical electrode and a thermistor.

68. The biosensor according to claim 56, in which the surface is operably associated with a member selected from the group consisting of a piezoelectric crystal, an electrochemical electrode and a thermistor.

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69. The biosensor according to claim 45, wherein said surface of the biosensor comprises gold.

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70. The biosensor according to claim 56, wherein said surface of the biosensor comprises gold.

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A method of using the biosensor according to claim 45 to determine the presence or amount of a protein, a virus or a cell, comprising the steps of:

exposing the biosensor to a sample containing a protein, a virus or a cell to be measured,

binding a protein, virus or cell to the biosensor, and measuring the presence or amount of the protein, virus or cell in the sample.

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72. A method of using the biosensor according to claim to determine the presence or amount of a protein, a virus or a cell, comprising the steps of:

exposing the biosensor to a sample containing a protein, a virus or a cell to be measured,

binding a protein, virus or cell to the biosensor, and measuring the presence or amount of the protein, virus or cell in the sample.

73. The biosensor according to claim 22, wherein:

the spacer molecule comprises an alkyl chain of the type $(-CH_2)_n$, in which n is an integer from 2 to 8.

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74. The biosensor according to claim 48, wherein:

the spacer molecule comprises an alkyl chain of the type $(-CH_2)_n$, in which n is an integer from 2 to 8.

56. The biosensor according to claim 59, wherein:

the spacer molecule comprises an alkyl chain of the type $(-CH_2)_n$, in which n is an integer from 2 to 8.

78. The biosensor according to claim 22, wherein:

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the spacer molecule comprises an aromatic group-containing structure; and the binding group comprises a member selected from the group consisting of -S-, -NH-CO-, -CO-NH-, -NH-, and -N=N-.

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M. The biosensor according to claim 48, wherein:

the spacer molecule comprises an aromatic group-containing structure; and the binding group comprises a member selected from the group consisting of -S-, -NH-CO-, -CO-NH-, -NH-, and -N=N-.

76. The biosensor according to claim 56, wherein:

the spacer molecule comprises an aromatic group-containing structure; and the binding group comprises a member selected from the group consisting of -S-, -NH-CO-, -CO-NH-, -NH-, and -N=N-.

79. The biosensor according to claim 22, wherein a chemical group is present between the surface and the binding group.

8%. The biosensor according to claim 7%, wherein the chemical group is a -CO-CH₂CH₂-S- group.

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